

# **Pediatric circumcision using n-butyl-cyanoacrylate plus MS monomer:**

## **bacteriostatic and cosmetic advantages over suture.**

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## **Abstract**

### **Purpose**

The aim of this study is to compare the use of n-butyl-cyanoacrylate plus MS monomer (NBCA-MS) glue with suture in pediatric circumcision.

### **Materials and Methods**

Between January 2013 and January 2017, 160 consecutive pediatric patients (range 18 months-14 years, mean 7.5 years) underwent circumcision. Eighty patients were randomly allocated to the glue group and 80 to the suture group. In the glue group wound margins were approximated by n-butyl-cyanoacrylate plus MS monomer (NBCA-MS) glue while in the suture group interrupted 5-0 polyglycolic acid stitches were used. Each group was randomized into two subgroups: forty patients (subgroup A) received antibiotics (6 days of Amoxicillin/clavulanic acid) and 40 (subgroup B)

patients did not receive any post-operative drug. Operative time, pain score, postoperative complications and cosmesis were statistically evaluated.

## Results

The operation time was significantly higher in the suture group; pain severity and duration were not significantly lower in the glue group. In the 80 patients of the glue group no complications occurred; in the suture group 20 (25%) cases of wound edema and 6 (7,5%) infections occurred. Cosmesis evaluation 1 and 6 months postoperatively showed significant better results in the glue group.

## Conclusion

Our study demonstrates that NBCA-MS glue wound closure in pediatric circumcision is a safe and effective technique.

## Introduction

The word "circumcision" comes from Latin *circum* (meaning "around") and *cœdere* (meaning "to cut"), and consists in removing the penis foreskin (prepuce). It has been practiced since the Egyptian times and it is still one of the most common surgical procedures: in the United States, nearly 75% of male population has been circumcised for ethnic, religious or medical reasons [1-3]. It has been widely proved that circumcised infants have a lower risk of urinary tract infections and

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balanoposthitis and seems to be a protective factor against sexual transmitted infections (STIs), such as HPV, HSV-2 and HIV [4-5]. Moreover, circumcision lowers the risk for penile cancer since predisposing conditions like phimosis, smegma, balanitis and HPV infection are uncommon [6]. Meanwhile, according to other authors there aren't randomized studies that could prove that rate of HWI could be reduced by circumcision[7]. The incidence and prevalence of most sexually transmitted diseases in Western Europe cannot be significantly influenced by circumcision, assuming regular genital hygiene [8].

Different techniques have been described in literature; the most used are the "Plastibell", the "sleeve resection", the "squeeze" and the "guillotine" technique [4].

Circumcision is a commonly performed operation and absorbable sutures have traditionally been used for closure. The two most common complications of circumcision are bleeding and wound infection [8], but also the penile adhesions after the surgery [9]. Male circumcision had a relatively low incidence of adverse events overall, especially if the procedure was performed during the first year of life, but rose 10–20 fold when performed after infancy [10].

The cyanoacrylate tissue glue has been claimed to have the advantages of being hemostatic [11], bacteriostatic [12-15] and easy to use; moreover, the stitch marks along the suture lines can be avoided, allowing for better cosmetic results.

The aim of our prospective randomized study is to show advantages of using n-butyl-cyanoacrylate plus MS monomer (NBCA-MS) tissue glue compared to absorbable sutures in pediatric circumcision, with particular focus on bacteriostatic effect.

## Methods

All the patients were fully informed about the study and informed consent was obtained. The study was conducted from January 2013 to January 2017. Totally, 160 boys of 18 months to 14 years of age (mean 7,5 years) underwent elective circumcision because of ethnical/religious reason in 125 cases and for congenital and acquired phimosis in 35.

80 patient were randomly assigned to the experimental group (glue group) and 80 patients to the control group (suture group). No patient underwent pre-operative treatment nor ointment therapy. Each group was further randomized into two subgroups: forty patients (subgroup A, mean age 72 years) received six days antibiotic therapy [9] and 40 (subgroup B, mean age 78 years) did not received any post-operative drug ([Figure 1](#)). In subgroup A, 57 intervention were for ritual purposes, while 23 for medical reasons (phimosis, paraphimosis, lichen, balanopostitis); in subgroup B 52 were rituals , while 28 medicals. The randomization timing we used was a mean of 12 days before the intervention (range 7-14 days)<sup>[16]</sup>

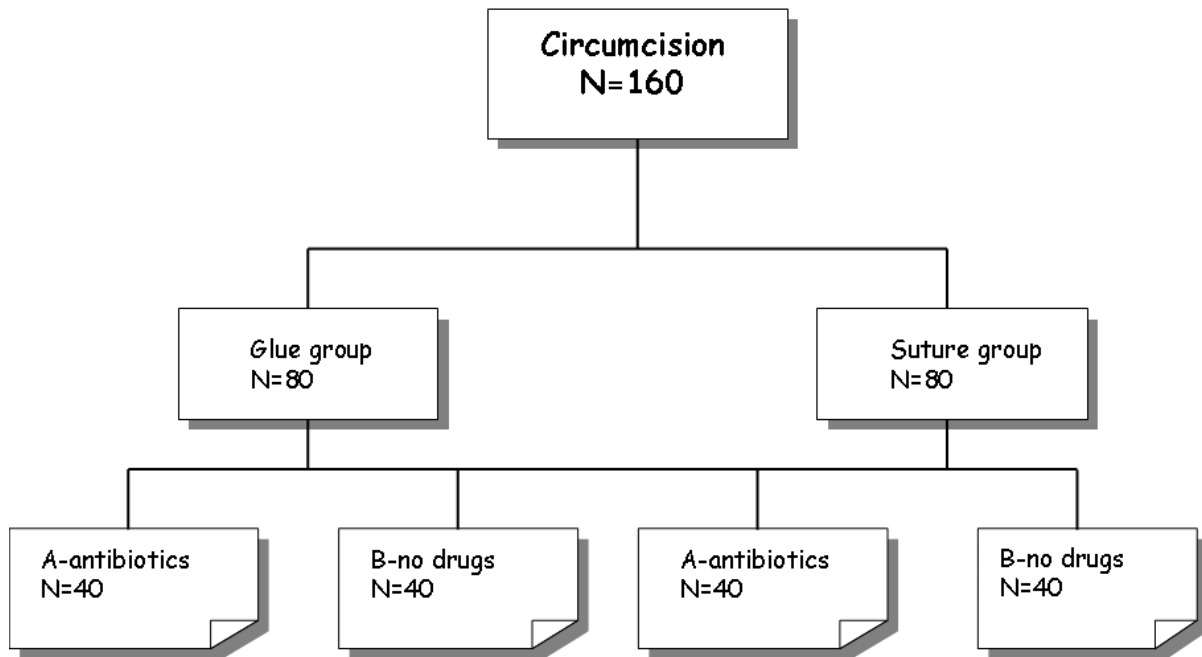


Figure 1

Study flowchart.

Patients were divided in four group and the following clinical events were searched: edema, bleeding, wound dehiscence and infection.

Postoperative antibiotic regimen included 6 days of amoxicillin/clavulanic acid (45mg/6,4mg/kg/day).

All patients were operated by the same surgeon under general anesthesia to avoid the bias. After povidone iodine disinfection, the penis foreskin was placed in traction and the skin was incised with the cold-scalpel surgery technique along an imaginary line starting from the frenulum and coming obliquely down following the corona glandis penis; the skin was then separated from dartos by round-tip scissors and excised with part of the mucosal cuff. Meticulous hemostasis was ensured by bipolar cauterization.

In the control group the mucocutaneous approximation was done by polyglycolic acid 5-0 stitches in the traditional manner. In the experimental group, a 0.25 ml dose of n-butyl-cyanoacrylate plus MS monomer (NBCA-MS) glue (Glubran2\*, GEM, Italy) was used for wound closure; we used a linear glue application, drying in about 1 minute (it was clear when the glue surface opacified), which peeled spontaneously off in around 10 days. A damp gauze was used to protect the glans and the urethra to avoid accidental contact with the glue ([Figure 2](#)). Mucocutaneous approximation was achieved by using a pair of toothed forceps holding the two edges of wound firmly together.

All patients were discharged the day after surgery with recommendation of daily iodine wound disinfection.

Post-operative pain was assessed after 4, 8 and 24 hours respectively, using Children and Infants Postoperative Pain Scale (ChIPPS) <sup>[15]</sup> .

On the 7<sup>th</sup> postoperative day after circumcision, wound was assessed for signs of infection (pain, swelling, fever, leucocytosis) bleeding and edema (highlighted by a swelling around, or just below the glandis); cosmetic outcomes were evaluated by a pediatric surgeon not participating the study, 30 and 180 days postoperatively.

Wound cosmesis was graded as poor, good or excellent, according to Visual Analog Scale (VAS) for scar evaluation [17].

Categorical data were described by absolute and relative frequency whereas quantitative data by mean and standard deviation. To analyze qualitative variables as “postoperative pain” and “clinical events” related to the groups, chi square test (or Fisher’s exact test when appropriate) and z-test for two proportions were applied. To compare continuous variables as “surgical case duration” and “time spent for sutureless circumcision” with the groups, t-test (two- tailed) was used.

A p-value less than 0.05 was considered statistically significant. All analyzes, descriptive and inferential, were performed by SPSS v.24 technology.



## Results

The comparison between the two groups shows a mean of surgical case duration of 20 minutes  $\pm$  1.5 (sd) when the suture were used whereas the mean of time spent for sutureless circumcision was 14 minutes  $\pm$  1.3 (sd). This difference was found as highly significant ( $p < 0.001$ ).

Regarding postoperative pain, there was no significant reduction when glue was used (Table I).

Table I

Postoperative pain (ChIPPS: Children and Infants Postoperative Pain scale) compared with groups.

	After 4 hours		After 8 hours		After 24 hours	
	0	1+2	0	1+2	0	1
<b>Score (ChIPPS)</b>						
<b>Suture group</b>	75	5	76	4	78	2
<b>Glue group</b>	79	1	80	0	80	0
<b><i>p-value</i></b>	0,212		0.129		0,477	

The comparison of complications using the cyanoacrylate and the suturing for wound approximation of circumcision is shown in Table II. Wound assessment on the 7th postoperative day revealed 20 cases (25%) of edema occurred in the suture group (11 cases , 27,5%, in subgroup A and 9 cases, 22,5%, in subgroup B) while no patient presented with wound edema in the glue group: this difference was statistically significant ( $p < 0.001$  and  $p < 0.005$ , in the comparisons between the two data subgroups). No patients in both groups, treatment and control, experienced wound bleeding or dehiscence. There are also significant differences between the percentage of infection in the suture-no drug group compared to the Glue-no drug group (15% compared to 0%;  $p = 0.034$ ), in fact six patients of the suture group, all previously randomized for not receiving postoperative antibiotics , showed serous wound discharge as a sign of infection, while conversely no infection occurred in the glue group.

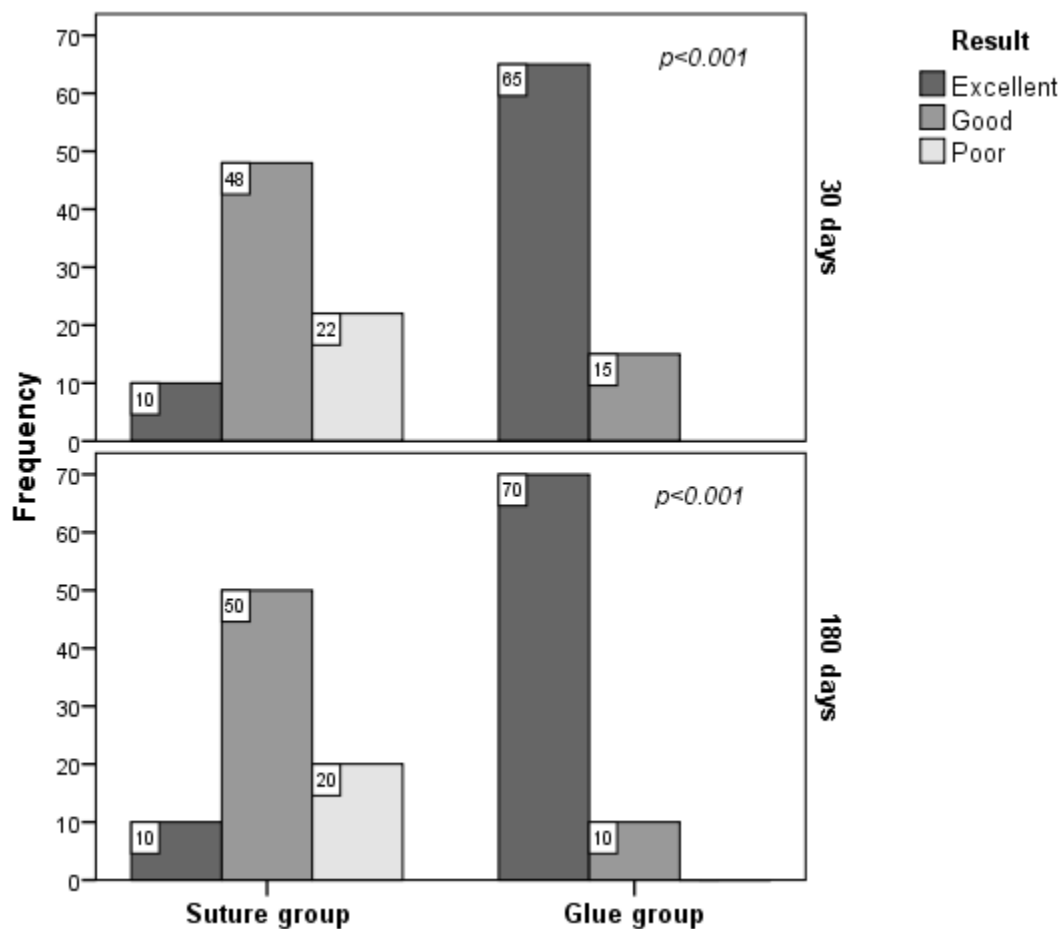
Table II  
Postoperative complications compared with groups.

Clinical events	Suture Group	Glue Group	p-value	Suture group	Glue group	p-value
	Drug	Drug		No drug	No drug	
<b>Edema</b>	11 (27.5%)	0 (0%)	<i>0.001</i>	9 (22.5%)	0 (0%)	<i>0.005</i>
<b>Bleeding</b>	0	0	-	0	0	-
<b>Wound dehiscence</b>	0	0	-	0	0	-
<b>Infection</b>	0	0	-	6 (15%)	0 (0%)	<i>0.034</i>

Thirty days after circumcision, the regularity of the wound edges and the overall cosmesis were better in the glue group; this difference was found to be highly significant ( $p < 0.001$ ). Similarly, after 180 days cosmetical advantage of tissue glue was confirmed (Table III). Moreover it is important to assess the fact that in our series the cost of intervention using the glue was cheaper than in the suture group; in fact 0,25ml of Glubran2\* (registered brand) cost 27US\$, against the 49,5US\$ of the suture).

Table III

Aesthetical results after 30 days and after 180 days compared with groups.



## Discussion

Traditionally, the circumcision wound is closed using interrupted absorbable suture; currently, alternative methods are being proposed to overcome the shortcomings of the conventional closure like complications, postoperative pain, time consumed and cosmetic problems; the use of glue seems to solve all these issues providing greater aesthetic results [18-20], also the the incidence of postoperative bleeding and

hematoma formation in pediatric circumcision is reduced with the use of tissue glue [21] .

Cyanoacrylate is a liquid-adhesive monomer which polymerizes on contact with basic substances such as blood or water. The polymerization occurs at room temperature and does not require the use of solvent or an added catalyst [22-25].

As already showed by Voznesensky et al, the use of glue can lead to a reduction in the operative time with beneficial economic effects [26]; similarly in our work the sutureless technique allowed to significantly shorten the time needed for surgery. Moreover, tissue glue has an important haemostatic effects [27]; in haemophilic patients the use of glue reduces the duration of clotting factor replacement after surgery, thus decreasing the high costs of treatment [28].

Finally, bacteriostatic advantage of n-butyl-cyanoacrylate has been already stated in the literature [29-30], and , to our knowledge, our study confirming the antimicrobial effects of NBCA-MS in a group of pediatric patients undergoing circumcision.

In the glue group, no infection occurred irrespectively of the postoperative antibiotic therapy was administered or not, demonstrating a protective role of the n-butyl-cyanoacrylate against infections.

Avoiding the use of antibiotics is considerably favorable in pediatric patients, who dislike the administration of drugs by mouth as well as the application of local therapies for wound care.

In addition, avoiding the use of antibiotics helps to prevent bacterial resistance and unexpected adverse reactions.

## Conclusion

In conclusion our study confirms that, in comparison with the traditional procedure, the sutureless technique with NBCA-MS for wound closure in pediatric circumcision allows to reduce the operative time, provides better cosmetic results and protects against infections.

**Conflict of Interest:** The authors declare that they have no conflict of interest.

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\* Glubran2 is a product of the Italian Pharmaceutical company GEM Italy. GEM Italy produces and commercializes medical devices since 1994.